U.S.S.N. 10/003,468 Attorney Docket No. FMI-001RCE2

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

1. (**Currently Amended**) A method for forming a two-dimensional ordered array of proteins,

comprising:

contacting a population of proteins with a gas-aqueous interface without using a

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detergent or solubilizing agent;

laterally compressing by planar membrane compression, said population to an appropriate pressure packing density, such that a two-dimensional ordered array of said proteins is formed at said interface, wherein said appropriate packing density is below a critical density

point proteins are not solubilized using detergent.

2-3. (Cancelled).

4. (**Previously Presented**) The method of claim 1, wherein said protein is a membrane

protein, a cellular receptor, an orphan receptor, receptor tyrosine kinase, an EPH receptor, an ion

channel, a cytokine receptor, an multisubunit immune recognition receptor, a chemokine

receptor, a growth factor receptor, or a G-protein coupled receptor.

5. (Previously Presented) The method of claim 1, wherein said protein is contacted with said

interface in the presence of lipids.

6. (**Previously Presented**) The method of claim 1, further comprising applying said proteins

to said interface in proteoliposomes, liposomes, or a cellular membrane.

7. (Cancelled).

8. (**Previously Presented**) The method of claim 1, wherein said interface is an air-aqueous

interface.

Claims 9-62 (Cancelled).

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63. (**Currently Amended**) A method for forming a two- or three-dimensional ordered array of water insoluble membrane proteins, comprising:

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contacting a population of water insoluble membrane proteins with a gas-aqueous interface without using a detergent or solubilizing agent, wherein said population of membrane proteins are applied to said interface in a proteoliposome;

laterally compressing <u>by planar membrane compression</u>, said population to an appropriate <u>pressure packing density</u>, such that a two- or three-dimensional ordered array of said water insoluble membrane proteins is formed at said gas-aqueous interface.

64. (**Currently Amended**) A method for forming a three-dimensional ordered array of water insoluble membrane proteins, comprising:

contacting a population of water insoluble membrane proteins with a gas-aqueous interface without using a detergent or solubilizing agent;

laterally compressing by planar membrane compression, said population to an appropriate pressure packing density, such that a three-dimensional ordered array of said water insoluble membrane proteins is formed at said interface, wherein said appropriate pressure packing density is above a critical density point for the formation of a two-dimensional ordered array of said water insoluble membrane proteins molecules.

Claims 65-66. (Cancelled).

- 67. (**Previously Presented**) The method of claim 1, wherein said two-dimensional ordered array is a two-dimensional crystalline array.
- 68. (**Previously Presented**) The method of claim 64, wherein said three-dimensional ordered array is a three-dimensional crystalline array.
- 69. (**Currently Amended**) The method of claim 1 [[3]], wherein said protein is a membrane protein, a cellular receptor, an orphan receptor, receptor tyrosine kinase, an EPH receptor, an ion channel, a cytokine receptor, an multisubunit immune recognition receptor, a chemokine receptor, a growth factor receptor, or a G-protein coupled receptor.
- 70. (**Currently Amended**) The method of claim 64 [[3]], wherein said water insoluble protein is contacted with said interface in the presence of lipids.

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71. (**Currently Amended**) The method of claim <u>64</u> [[3]], further comprising applying said

water insoluble proteins to said interface in proteoliposomes, liposomes, or a cellular membrane.

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Claims 72-73 (Cancelled).

74. (**Currently Amended**) A method for forming a two- or three- dimensional ordered array of <u>water insoluble membrane</u> proteins suitable for use in crystallography to determine said <u>protein's water insoluble membrane proteins'</u> structure, comprising:

contacting a population of <u>water insoluble membrane</u> proteins with a gas-aqueous interface <u>without using a detergent or solubilizing agent;</u>

laterally compressing <u>by planar membrane compression</u>, said population to an appropriate <u>pressure packing density</u>, such that a two- <u>or three-</u> dimensional ordered array of said <u>water insoluble membrane proteins</u> is formed at said interface, wherein the structure of said <u>water insoluble membrane proteins</u> using said two- or three- dimensional ordered array can be determined to a resolution of 5 Å or higher.

75-76. (Canceled)

- 77. (New) The method of claim 74 wherein said ordered array is formed in the absence of a ligand of said water insoluble membrane protein.
- 78. (New) The method of claim 77 wherein said appropriate packing density is below a critical density point such that a two dimensional ordered array is formed at said interface.
- 79. (New) The method of claim 77 wherein said appropriate packing density is above a critical density point such that a three dimensional ordered array is formed at said interface.
- 80. (New) The method of claim 77 further comprising applying said water insoluble proteins to said interface in proteoliposomes.
- 81. (New) The method of claim 80 wherein said water insoluble membrane proteins in said ordered array maintain orientation in same direction.
- 82. (New) The method of claim 81 further comprising spreading said lysed proteoliposomes at said interface to form a planar lipid-protein film.

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83. (New) The method of claim 82 further comprising achieving an equilibrium pressure between said lipid-protein film and unlysed proteoliposomes.

- 84. (New) The method of claim 83 wherein said equilibrium pressure is in the range of 20 to 38 mN/m.
- 85. (New) The method of claim 84 further comprising compressing said lipid-protein film from 40 cm² to 11 cm².
- 86. (New) The method of claim 85 further comprising compressing said lipid-protein film at a rate of 500 mm²/min.
- 87. (New) The method of claim 86 further comprising compressing said lipid-protein film to a density corresponding to a pressure between 35 to 45 mN/m.
- 88. (New) The method of claim 74 wherein said ordered array is formed in the presence of a ligand of said water insoluble membrane protein.
- 89. (New) A method for forming a two- or three- dimensional ordered array of water insoluble membrane proteins suitable for use in crystallography to determine said water insoluble proteins' structure, comprising:

contacting a population of water insoluble membrane proteins with a gas-aqueous interface in proteoliposomes without using a detergent or solubilizing agent;

lysing said proteoliposomes;

spreading said lysed proteoliposomes at said interface to form a planar lipid-protein film;

achieving an equilibrium pressure in the range of 20 to 38 mN/m between said lipid-protein film and unlysed proteoliposomes;

laterally compressing by planar membrane compression, said lipid-protein film from 40 cm² to 11 cm² to a density corresponding to a pressure between 35 to 45 mN/m, such that a two- or three- dimensional ordered array of said water insoluble membrane proteins is formed at said interface, wherein the structure of said water insoluble membrane proteins using said two- or three- dimensional ordered array can be determined to a resolution of 5 Å or higher.